IDW 18/CC

Sterne Kessler Goldstein Fox
ATTORNEYS AT LAW

January 18, 2006

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Commissioner for Patents
PO Box 1450
Alexandria, VA 22313-1450

Attn: Mail Stop Amendment

Re: U.S. Utility Patent Application
Application No. 10/698,928; Filed: October 31, 2003
For: **Novel Cytarabine Monophosphate Prodrugs**
Inventors: Boyer *et al.*
Our Ref: 2358.0180002/RWE/AES

Sir:

Transmitted herewith for appropriate action are the following documents:

1. Credit Card Payment Form (PTO-2038) in the amount of **$180.00** to cover Information Disclosure Statement Filing fee (37 C.F.R. § 1.17(p));

2. First Supplemental Information Disclosure Statement Filing Under 37 C.F.R. § 1.97(c);

3. One (1) page of Form PTO/SB/08A citing seventeen (17) documents;

4. Copies of nine (9) of the cited documents (Documents FE-FM);

5. Ten (10) pages of Form PTO/SB/08B citing ninety-nine (99) documents;

6. Copies of the ninety-nine (99) cited documents (Document FN-JH); and

7. One (1) return postcard.

It is respectfully requested that the attached postcard be stamped with the date of filing of these documents, and that it be returned to our courier. In the event that extensions of time are necessary to prevent abandonment of this patent application, then such extensions of time are hereby petitioned.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.

Ann E. Summerfield
Attorney for Applicants
Registration No. 47,982

AES/law
Enclosures

484280_1.DOC

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:	Confirmation No.: 2990
BOYER *et al.*	Art Unit: 1626
Appl. No.: 10/698,928	Examiner: T. Solola
Filed: October 31, 2003	Atty. Docket: 2358.0180002/RWE/AES
For: **Novel Cytarabine Monophosphate Prodrugs**	

First Supplemental Information Disclosure Statement Filing Under 37 C.F.R. § 1.97(c)

Mail Stop Amendment

Commissioner for Patents
PO Box 1450
Alexandria, VA 22313-1450

Sir:

Listed on accompanying Forms PTO/SB/08A and PTO/SB/08B are documents that may be considered material to the examination of this application, in compliance with the duty of disclosure requirements of 37 C.F.R. §§ 1.56, 1.97 and 1.98. The numbering on this First Supplemental Information Disclosure Statement is a continuation of the numbering in Applicants' Information Disclosure Statement filed on February 26, 2004 in connection with the above-captioned application.

Copies of documents **FE-FM and FN-JH** are submitted herewith. However, in accordance with 37 C.F.R. § 1.98(a)(2), copies of U.S. patents and patent application publications, documents **EW-FD**, cited on the attached Form PTO/SB/08A are not submitted.

In accordance with 37 C.F.R. § 1.98(a)(3), Applicants' undersigned representative submits the following discussion of the relevance of the non-English language document **IO** cited on Form PTO/SB/08B:

Document **IO,** "SARTILLO-PISCIL, F., *et al.*, "Fosfato-ésteres cíclicos diastereoisoméricos: 5-bromo-4-fenil-2-fenoxi-2-oxo-1,3,2-dioxafosforinanos, precursores de radicales libres alquilo β-fosfatoxi y generadores de radicales catiónicos en medio no oxidativo," *Revista de la Sociedad Química de México 46*:330-334, Sociedad Quimica de Mexico (2002), is in the Spanish language. An English language abstract of document **IO** is located on the face page of said document.

The Examiner's attention is directed to the following co-pending U.S. Patent Application, which is directed to related technical subject matter:

Application No. 10/698,924, inventors Reddy, K.R., *et al.*, filed October 31, 2003, published as 2004/0192651 A1, cited herein as document **FC**, and

Application No. 11/145,194, inventors Erion, M.D., *et al.*, filed June 3, 2005, published as US 2005/0288240 A1, cited herein as document **FD.**

In accordance with the recent Federal Circuit decision in *Dayco Prods., Inc. v. Total Containment, Inc.* 329 F.3d 1358 (Fed. Cir. 2003), Applicants submit herewith an Office Action, which is also directed to related technical subject matter:

Office Action for related U.S. Patent Application No. 10/698,924, inventors Reddy, K.R., *et al.*, filed October 31, 2003, mailed June 22, 2005, cited herein as document **JH.**

The identification of this U.S. Patent Application and Office Action is not to be construed as a waiver of secrecy as to that application now or upon issuance of the

present application as a patent. The Examiner is respectfully requested to consider the cited applications and the art cited therein during examination.

Where the publication date of a listed document does not provide a month of publication, the year of publication of the listed document is sufficiently earlier or later than the effective U.S. filing date and any foreign priority date so that the month of publication is not in issue. Applicants have listed publication dates on the attached IDS Forms based on information presently available to the undersigned. However, the listed publication dates should not be construed as an admission that the information was actually published on the date indicated.

Applicants reserve the right to establish the patentability of the claimed invention over any of the information provided herewith, and/or to prove that this information may not be prior art, and/or to prove that this information may not be enabling for the teachings purportedly offered.

This statement should not be construed as a representation that a search has been made, or that information more material to the examination of the present patent application does not exist. The Examiner is specifically requested not to rely solely on the material submitted herewith.

This First Supplemental Information Disclosure Statement is being filed more than three months after the U.S. filing date AND after the mailing date of the first Office Action on the merits, but before the mailing date of a Final Rejection, or Notice of Allowance, or an action that otherwise closes prosecution in the application. Attached is

our PTO-2038 Credit Card Payment Form in the amount of $180.00 in payment of the fee under 37 C.F.R. § 1.17(p).

It is respectfully requested that the Examiner initial and return copies of the enclosed IDS Forms, and indicate in the official file wrapper of this patent application that the documents have been considered.

The U.S. Patent and Trademark Office is hereby authorized to charge any fee deficiency, or credit any overpayment, to our Deposit Account No. 19-0036.

Respectfully submitted,

STERNE, KESSLER, GOLDSTEIN & FOX P.L.L.C.

Ann E. Summerfield

Ann E. Summerfield
Attorney for Applicants
Registration No. 47,982

Date: January 18, 2006

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Substitute for form 1449/PTO

FIRST SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(Use as many sheets as necessary)

Sheet 1 of 1

Complete if Known	
Application Number	10/698,928
Filing Date	October 31, 2003
First Named Inventor	Serge Boyer
Art Unit	1626
Examiner Name	Solola, Taofiq A.
Attorney Docket Number	2358.0180002/RWE/AES

U.S. PATENT DOCUMENTS

Examiner Initials*	Cite No.[1]	Document Number Number-Kind Code[2] (If Known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
	EW	3,018,302	01/23/1962	Bielefeld *et al.*	
	EX	4,952,740	08/28/1990	Juge *et al.*	
	EY	5,663,159	09/02/1997	Starrett Jr., *et al.*	
	EZ	6,752,981	06/22/2004	Erion *et al.*	
	FA	2003/0225277 A1	12/04/2003	Kopcho *et al.*	
	FB	2003/0229225 A1	12/11/2003	Reddy *et al.*	
	FC	2004/0192651 A1	09/30/2004	Reddy *et al.*	
	FD	2005/0288240 A1	12/29/2005	Erion *et al.*	

FOREIGN PATENT DOCUMENTS

Examiner Initials*	Cite No.[1]	Foreign Patent Document Country Code[3] Number[4] Kind Code[5] (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T[6]
	FE	EP 0161955 A1	11/21/1985	Merck & Co., Inc.		
	FF	EP 0180276 A1	05/07/1986	Oce-Andeno B.V.		
	FG	EP 0338372 A2	10/25/1989	American Cyanamid Co.		
	FH	EP 0353692 B1	10/04/1995	Nissan Chemical Ind., Ltd.		
	FI	EP 0481214 B1	06/24/1998	Inst. Organic Chem. & Biochem. Acad. Sci. Czech. Repub.		
	FJ	WO 91/19721 A1	12/26/1991	Glazier		
	FK	WO 96/01267 A1	01/18/1996	Takeda Chemical Ind., Ltd.		
	FL	WO 97/03679 A1	02/06/1997	Cephalon, Inc.		
	FM	WO 00/52015 A2	09/08/2000	Metabasis Therapeutics, Inc.		

Examiner Signature		Date Considered	

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This collection of information is required by 37 CFR 1.97 and 1.98. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 2 hours to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. **SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.**

If you need assistance in completing the form, call 1-800-PTO-9199 (1-800-786-9199) and select option 2.

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Substitute for form 1449/PTO				*Complete if Known*	
FIRST SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT BY APPLICANT *(Use as many sheets as necessary)*				Application Number	10/698,928
				Filing Date	October 31, 2003
				First Named Inventor	Serge Boyer
				Art Unit	1626
				Examiner Name	Solola, Taofiq A.
Sheet	1	of	10	Attorney Docket Number	2358.0180002/RWE/AES

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No.[1]	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume issue number(s), publisher, city and/or country where published	T[2]
	FN	ALEXANDER, P., *et al.*, "Preparation of 9-(2-Phosphonomethoxyethyl) Adenine Esters as Potential Prodrugs," *Collect. Czech. Chem. Commun.* 59:1853-1869, Czech Academy of Sciences, Institute of Organic Chemistry and Biochemistry (1994)	
	FO	AMIN, D., *et al.*, "1-Hydroxy-3-(methylpentylamino)-propylidene-1,1-bisphosphonic Acid as a Potent Inhibitor of Squalene Synthase," *Arzneim.-Forsch/Drug Res. 46*:759-762, Blackwell Publishing, Inc. (1996)	
	FP	ATIQ, O., *et al.*, "Treatment of Unresectable Primary Liver Cancer with Intrahepatic Fluorodeoxyuridine and Mitomycin C Through an Implantable Pump," *Cancer 69*:920-924, John Wiley and Sons, Inc. (1992)	
	FQ	AUBERSON, Y., *et al.*, "N-Phosphonoalkyl-5-Aminomethylquinoxaline-2,3-Diones: *In Vivo* Active AMPA and NMDA-(Glycine) Antagonists," *Bioorg. Med. Chem. Lett. 9*:249-254, Elsevier Science Ltd. (1999)	
	FR	BALTHAZOR, T. and Grabiak, R.C., "Nickel-Catalyzed Arbuzov Reaction: Mechanistic Observations," *J. Org. Chem. 45*:5425-5426, American Chemical Society (1980)	
	FS	BEAUCAGE, S.L. and Iyer, R.P., "The Synthesis of Modified Oligonucleotides by the Phosphoramidite Approach and Their Applications," *Tetrahedron 49*:6123-6194, Pergamon Press Ltd. (1993)	
	FT	BESPALOV, A., *et al.*, "Prolongation of morphine analgesia by competitive NMDA receptor antagonist D-CPPene (SDZ EAA 494) in rats," *Eur. J. Pharmacol. 351*:299-305, Elsevier Science B.V. (1998)	
	FU	BIJSTERBOSCH, M., *et al.*, "Disposition of the Acyclic Nucleoside Phosphonate (S)-9-(3-Hydroxy-2-Phosphonylmethoxyproply)Adenine," *Antimicrob. Agents Chemother. 42*:1146-1150, American Society for Microbiology (1998)	
	FV	BIRD, J., *et al.*, "Synthesis of Novel *N*-Phosphonoalkyl Dipeptide Inhibitors of Human Collagenase," *J. Med. Chem. 37*:158-169, American Chemical Society (1994)	
	FW	BORCH, R.F. and Millard, J.A., "The Mechanism of Activation of 4-Hydroxycyclophosphamide," *J. Med. Chem. 30*:427-431, American Chemical Society (1987)	

Examiner Signature		Date Considered	

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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Sheet	2	of	10	Attorney Docket Number	2358.0180002/RWE/AES

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	FX	BRILL, T. and Landon, S.J., "Arbuzov-like Dealkylation Reactions of Transition-Metal-Phosphite Complexes," *Chem. Rev. 84*:577-585, American Chemical Society (1984)	
	FY	CAMPAGNE, J.-M., *et al.*, "Synthesis of Mixed Phosphate Diester Analogues of Dipeptides using BOP or PyBOP Reagents," *Tetrahedron Lett.* 34:6743-6744, Pergamon Press Ltd. (1993)	
	FZ	CAMPBELL, D.A., "The Synthesis of Phosphonate Esters, an Extension of the Mitsunobu Reation," *J. Org. Chem.* 57:6331-6335, American Chemical Society (1992)	
	GA	CASARA, P., *et al.*, "Synthesis of Acid Stable 5'-O-Fluoromethyl Phosphonates of Nucleosides. Evaluation as Inhibitors of Reverse Transcriptase," *Bioorg. Med. Chem. Lett.* 2:145-148, Pergamon Press plc (1992)	
	GB	CASTEEL, D. and Peri, S.P., "Steric and Electronic Effects in the Aryl Phosphate to Arylphosphonate Rearrangement," *Synthesis* (9):691-693, Georg Thieme Verlag KG (1991)	
	GC	CHEN, L. and Waxman, D.J., "Intratumoral Activation and Enhanced Chemotherapeutic Effect of Oxazaphosphorines following Cytochrome P-450 Gene Transfer: Development of a Combined Chemotherapy/Cancer Gene Therapy Strategy," *Cancer Res. 55*:581-589, The American Association for Cancer Research (1995)	
	GD	CHEN, L., *et al.*, "Sensitization of Human Breast Cancer Cells to Cyclophosphamide and Ifosfamide by Transfer of a Liver Cytochrome P450 Gene," *Cancer Res. 56*:1331-1340, The American Association for Cancer Research (1996)	
	GE	COOPER, D.B., *et al.*, "Use of Carbohydrate Derivatives for Studies of Phosphorus Stereo-chemistry. Part II. Synthesis and Configurational Assignments of 1,-3,2-Oxathiaphosphrinan-2-ones and 1,3,2-Dioxaphosphorinan-2-thiones," *J .Chem. Soc. Perkin I* 3/2422:1049-1052, Royal Society of Chemistry (1974)	
	GF	DEARFIELD, K., *et al.*, "Analysis of the genotoxicity of nine acrylate/methacrylate compounds in L5178Y mouse lymphoma cells," *Mutagenesis 4*:381-393, Oxford University Press (1989)	
	GG	DE CLERCQ, E., *et al.*, "A novel selective broad-spectrum anti-DNA virus agent," *Nature 323*:464-467, Nature Publishing Group (1986)	

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FIRST SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT BY APPLICANT

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Sheet 3 of 10

Complete if Known	
Application Number	10/698,928
Filing Date	October 31, 2003
First Named Inventor	Serge Boyer
Art Unit	1626
Examiner Name	Solola, Taofiq A.
Attorney Docket Number	2358.0180002/RWE/AES

NON PATENT LITERATURE DOCUMENTS

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	GH	DE LOMBAERT, S., *et al.*, "Pharmacological Profile of a Non-Peptidic Dual Inhibitor of Neutral Endopeptidase 24.11 and Endothelin-Converting Enzyme," *Biochem. Biophys. Res. Commun. 204*:407-412, Academic Press, Inc. (1994)	
	GI	DE LOMBAERT, S., *et al.*, "*N*-Phosphonomethyl Dipeptides and Their Phosphonate Prodrugs, a New Generation of Neutral Endopeptidase (NEP, EC 3.4.24.11) Inhibitors," *J. Med. Chem. 37*:498-511, American Chemical Society (1994)	
	GJ	DESOS, P., *et al.*, "Structure-Activity Relationships in a Series of 2(1*H*)-Quinolones Bearing Different Acidic Function in the 3-Position: 6,7-Dichloro-2(1*H*)-oxoquinoline-3-phosphonic Acid, a New Potent and Selective AMPA/Kainate Antagonist with Neuroprotective Properties," *J. Med. Chem. 39*:197-206, American Chemical Society (1996)	
	GK	DICKSON, J.K., *et al.*, "Orally Active Squalene Synthase Inhibitors: Bis((acyloxy)alkyl) Prodrugs of the α-Phosphonosulfonic Acid Moiety," *J. Med. Chem. 39*:661-664, American Chemical Society (1996)	
	GL	EDMUNDSON, R.S., *et al.*, "Cyclic Organophosphorous Compounds. Part 23. Configurational Assignments in the 4-Phenyl-1,3,2λ-dioxaphosphorinane Series. *X*-Ray Molecular Structure of cis-2-Benzylamino-4-phenyl-1,3,2-dioxaphosphorinane 2-Oxide," *J. Chem. Research (S)*, 122-123, Science Reviews Ltd. (1989)	
	GM	ENRIQUEZ, P., *et al.*, "Conjugation of Adenine Arabinoside 5'-Monophosphate to Arabinogalactan: Synthesis, Characterization, and Antiviral Activity," *Bioconjugate Chem. 6*:195-202, American Chemical Society (1995)	
	GN	ERION, M., *et al.*, "Design, Synthesis, and Characterization of a Series of Cytochrome P_{450} 3A-Activated Prodrugs (HepDirect Prodrugs) Useful for Targeting Phosph(on)ate-Based Drugs to the Liver," *J. Am. Chem. Soc. 126*:5154-5163, American Chemical Society (April 2004)	
	GO	ERION, M., *et al.*, "HepDirect™ Prodrugs: A Novel Strategy for Targeting Drugs to the Liver," *Hepatology 36*:301A, AASLD Abstract No. 551, John Wiley & Sons, Inc. (October 2002)	
	GP	ERION, M., *et al.*, "Liver-Targeted Drug Delivery Using HepDirect Prodrugs" *J. Pharmacol. Exper. Ther. 312*:554-560, American Society for Pharmacology and Experimental Therapeutics (February 2005)	
	GQ	ERION, M., "Liver-Targeted Nucleoside Prodrugs," presented at the *Gordon Research Conference: Purines, Pyrimidines and Related Substances*, Newport, RI, 38 pages (June-July 2003)	

Examiner Signature		Date Considered	

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				Filing Date	October 31, 2003
				First Named Inventor	Serge Boyer
				Art Unit	1626
				Examiner Name	Solola, Taofiq A.
Sheet	4	of	10	Attorney Docket Number	2358.0180002/RWE/AES

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No.[1]	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume number, publisher, city and/or country where published	T[2]
	GR	FARQUHAR, D., *et al.*, "Biologically-Cleavable Phosphate Protective Groups: 4-Acyloxy-1,3,2-Dioxaphosphorinanes as Neutral Latent Precursors of Dianionic Phosphates," *Tetrahedron Lett. 36*:655-658, Elsevier Science Ltd. (1995)	
	GS	FARQUHAR, D., *et al.*, "Biologically Reversible Phosphate-Protective Groups," *J. Pharm. Sci.* 72:324-325, American Chemical Society (1983)	
	GT	FARQUHAR, D., *et al.*, "5'-4-(Pivaloyloxy)-1,3,2-dioxaphosphorinan-2-yl]-2'-deoxy-5-fluorouridine: A Membrane-Permeating Prodrug of 5-Fluoro-2'-deoxyuridylic Acid (FdUMP)," *J. Med. Chem.* 38:488-495, American Chemical Society (1995)	
	GU	FARQUHAR, D., *et al.*, "Synthesis and Antitumor Evaluation of Bis[(pivaloyloxy) methyl] 2'-Deoxy-5-fluorouridine 5'-Monophosphate (FdUMP): A Strategy to Introduce Nucleotides into Cells," *J. Med. Chem. 37*:3902-3909, American Chemical Society (1994)	
	GV	FARQUHAR, D., *et al.*, Synthesis and Biological Evaluation of 9-[5'-(2-Oxo-1,3,2-oxazaphosphorinan-2-yl)-β-D-arabinosyl]adenine and 9-[5'-(2-Oxo-1,3,2-dioxaphosphorinan-2-yl)-β-D-arabinosyl]adenine: Potential Neutral Precursors of 9-[β-D-Arabinofuranosyl]adenine 5'-Monophosphate," *J. Med. Chem. 28*:1358-1361, American Chemical Society (1985)	
	GW	FARQUHAR, D., *et al.*, "Synthesis and Biological Evaluation of Neutral Derivatives of 5-Fluoro-2'-deoxyuridine 5'-Phosphate," *J. Med. Chem. 26*:1153-1158, American Chemical Society (1983)	
	GX	FIUME, L., *et al.*, "Inhibition of Hepatitis B Virus Replication by Vidarabine Monophosphate Conjugated with Lactosaminated Serum Albumin," *The Lancet* 2:13-15, The Lancet Publishing Group (1988)	
	GY	FREED, J.J., *et al.*, "Evidence for Acyloxymethyl Esters of Pyrimidine, 5'-Deoxyribonucleotides as Extracellular Sources of Active 5'-Deoxyribonucleotides in Cultured Cells," *Biochem. Pharm. 38*:3193-3198, Elsevier Inc. (1989)	
	GZ	FRIIS, G.J. and Bundgaard, H., "Prodrugs of phosphates and phosphonates: Novel lipophilic α-acyloxyalkyl ester derivatives of phosphate- or phosphonate containing drugs masking the negative charges of these groups," *Eur. J. Pharm. Sci. 4*:49-59, Elsevier Science B.V. (1996)	
	HA	GUIDA, W.C., *et al.*, "Structure-Based Design of Inhibitors of Purine Nucleoside Phosphorylase. 4. A Study of Phosphate Mimics," *J. Med. Chem. 37*:1109-1114, American Chemical Society (1994)	

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Sheet	5	of	10	Attorney Docket Number	2358.0180002/RWE/AES

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	HB	HE, K., *et al.*, "Inactivation of Cytochrome P450 3A4 by Bergamottin, a Component of Grapefruit Juice," *Chem. Res. Toxicol. 11*:252-259, American Chemical Society (1998)	
	HC	HILLERS, S., *et al.*, "Analogs of pyrimidinemono-and polynucleotides. VI. Phosphates of 1-(1,4-dihydroxy-2-pentyl)thymine and 1-(1,3-dihydroxy-2-propyl) uracil," *Chemical Abstracts* 89(17), Chemical Abstracts Service (1978)	
	HD	HIRAYAMA, N., *et al.*, "Structure and conformation of a novel inhibitor of angiotensin I converting enzyme - a tripeptide containing phosphonic acid," *Int. J. Pept. Protein Res. 38*:20-24, Blackwell Publishing (1991)	
	HE	HUNSTON, R., *et al.*, "Synthesis and Biological Properties of Some Cyclic Phosphotriesters Derived from 2'-Deoxy-5-fluorouridine," *J. Med. Chem. 27*:440-444, American Chemical Society (1984)	
	HF	KEENAN, R., *et al.*, "Pathology Reevaluation of the Kociba et al. (1978) Bioassay of 2,3,7,8-TCDD: Implications for Risk Assessment," *J. Tox. Envir. Health 34*:279-296 , Hemisphere Publishing Corporation (1991)	
	HG	KELLEY, J.L., *et al.*, "[[(Guaninylalkyl)phosphinico]methyl]phosphonic Acids. Multisubstrate Analogue Inhibitors of Human Erythrocyte Purine Nucleoside Phosphorylase," *J. Med. Chem. 38*:1005-1014, American Chemical Society (1995)	
	HH	KHAMNEI, S. and Torrence, P.F., "Neighboring Group Catalysis in the Design of Nucleotide Prodrugs," *J. Med. Chem. 39*:4109-4115, American Chemical Society (1996)	
	HI	KHORANA, H.G., *et al.*, "Cyclic Phosphates. III. Some General Observations on the Formation and Properties of Five-,Six- and Seven-membered Cyclic Phosphate Esters," J. Am. Chem. Soc. 79:430-436, American Chemical Society (1957)	
	HJ	KORBA, B.A., *et al.*, "Liver-Targeted Antiviral Nucleosides: Enhanced Antiviral Activity of Phosphatidyl-Dideoxyguanosine Versus Dideoxyguanosine in Woodchuck Hepatitis Virus Infection *In Vivo*," *Hepatology* 23:958-963, John Wiley & Sons, Inc. (1996)	
	HK	KRYUCHKOV, A.A., *et al.*, "Influence of Solvent on the Strength of Cyclic Oxygen-Containing Phosphorus Acids," *Bull. Acad. Sci. USSR, A translation of Izvestiya Akademii Nauk SSSR, Ser. Khim.* 36:1145-1148, Consultants Bureau (1987)	

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				Filing Date	October 31, 2003
				First Named Inventor	Serge Boyer
				Art Unit	1626
				Examiner Name	Solola, Taofiq A.
Sheet	6	of	10	Attorney Docket Number	2358.0180002/RWE/AES

NON PATENT LITERATURE DOCUMENTS			
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	HL	LEFEBVRE, I., *et al.*, "Mononucleoside Phosphotriester Derivatives with *S*-Acyl-2-thioethyl Bioreversible Phosphate-Protecting Groups: Intracellular Delivery of 3'-Azido-2',3'-dideoxythymidine 5'-Monophosphate," *J. Med. Chem.* 38:3941-3950, American Chemical Society (1995)	
	HM	LOK, A.S.F., *et al.*, "Neurotoxicity associated with adenine arabinoside monophosphate in the treatment of chronic hepatitis B virus infection," *J. Antimicrob. Chemotherap.* 14:93-99, Oxford University Press (1984)	
	HN	LU, X. and Zhu, J., "Palladium-Catalyzed Reaction of Aryl Polyfluoroalkanesulfonates with *O,O*-Dialkyl Phosphonates," *Synthesis (8)*:726-727, Georg Thieme Verlag (1987)	
	HO	LUDEMAN, S.M., *et al.*, "Synthesis and Antitumor Activity of Cyclophosphamide Analogues. 4. Preparation, Kinetic Studies, and Anticancer Screening of "Phenylketophosphamide" and Similar Compounds Related to the Cyclophosphamide Metabolite Aldophosphamide," *J. Med. Chem. 29*:716-727, American Chemical Society (1986)	
	HP	MACKENNA, D., *et al.*, "MB07133: A HepDirect™ Prodrug of Cytarabine Monophosphate for Use in Hepatocellular Carcinoma," *Heptaology 38(Suppl. 1)*:411A, AASLD Abstract No. 524, John Wiley & Sons, Inc. (October 2003)	
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	HS	MEIER, C., *et al.*, "Cyclic Saligenyl Phosphotriesters of 2',3'-Dideoxy-2',3'-didehydrothymidine (d4T) - A New Pro-Nucleotide Approach -" *Bioorg. Med. Chem. Lett.* 7:99-104, Elsevier Science Ltd. (1997)	
	HT	MEIJER, D.K.F. and van der Sluijs, P., "Covalent and Noncovalent Protein Binding of Drugs: Implications for Hepatic Clearance, Storage, and Cell-Specific Drug Delivery," *Pharm. Res.* 6:105-118, Plenum Publishing Corporation (1989)	
	HU	MELVIN, L.S., "An Efficient Synthesis of 2-Hydroxyphenylphosphonates" *Tetrahedron Lett.* 22:3375-3376, Pergamon Press Ltd. (1981)	

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Sheet 7 of 10

Complete if Known	
Application Number	10/698,928
Filing Date	October 31, 2003
First Named Inventor	Serge Boyer
Art Unit	1626
Examiner Name	Solola, Taofiq A.
Attorney Docket Number	2358.0180002/RWE/AES

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	HV	MEYER, R., *et al.*, "2'-*O*-Acyl-6-thioinosine Cyclic 3',5'-Phosphates as Prodrugs of Thioinosinic Acid," *J. Med. Chem. 22*:811-815, American Chemical Society (1979)	
	HW	MITCHELL, A., *et al.*, "Bioreversible Protection for the Phospho Group: Bioactivation of the Di(4-acyloxybenzyl) and Mono(4-acyloxybenzyl) Phosphoesters of Methylphosphonate and Phosphonacetate," *J. Chem. Soc. Perkin Trans. 1*, 2345-2353, Royal Society of Chemistry (1992)	
	HX	MITSUNOBU, O., "The Use of Diethyl Azodicarboxylate and Triphenylphosphine in Synthesis and Transformation of Natural Products," *Synthesis (1)*:1-28, Georg Thieme Verlag (1981)	
	HY	MONTAG, A., *et al.*, "The Effect of Dexamethasone Treatment on CYP3A Activity Distribution, the Liver Targeting of MB07133 and CYP3A Activity in a Highly Proliferating State in Rats," *Hepatology 40(Suppl. 1)*:649A, AASLD Abstract No. 1123, John Wiley & Sons, Inc. (2004)	
	HZ	MOORE, M., *et al.*, "Comparison of mutagenicity results for nine compounds evaluated at the *hgprt* locus in the standard and suspension CHO assays," *Mutagenesis 6*:77-85, Oxford University Press (1991)	
	IA	MURRAY, G., *et al.*, "Cytochrome P450 CYP3A in human renal cell cancer," *Brit. J. Cancer 79*:1836-1842, Nature Publishing Group (1999)	
	IB	MURRAY, G., *et al.*, "Cytochrome P450 Expression Is a Common Molecular Event in Soft Tissue Sarcomas," *J. Pathology 171*:49-52, John Wiley & Sons, Ltd. (1993)	
	IC	NAKAYAMA, K. and Thompson, W.J., "A Highly Enantioselective Synthesis of Phosphate Triesters," *J. Am. Chem. Soc. 112*:6936-6942, American Chemical Society (1990)	
	ID	NEIDLEIN, R., *et al.*, "Mild Preparation of 1-Benzyloxyiminoalkylphosphonic Dichlorides: Application to the Synthesis of Cyclic Phosphonic Deisters and Cyclic Monoester Amides," *Heterocycles 35*:1185-1203, Elsevier Science (1993)	
	IE	NIFANTYEV, E.E., *et al.*, "Synthesis and Structure of Some Stable Phospholane-Phospholanes," *Phosphorus, Sulfur and Silicon 113*:1-13, Overseas Publishers Association (1996)	

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	IF	OGG, M., *et al.*, "A reporter gene assay to assess the molecular mechanisms of xenobiotic-dependent induction of the human CYP3A4 gene *in vitro*," *Xenobiotica 29*:269-279, Taylor & Francis Ltd. (1999)	
	IG	OHASHI, K., *et al.*, "Synthesis of Phosphonosphingoglycolipid Found in Marine Snail Turbo Cornutus," *Tetrahedron Lett. 29*:1189-1192, Pergamon Press plc (1988)	
	IH	PETRAKIS, K. and Nagabhushan, T.L., "Palladium-Catalyzed Substitutions of Triflates Derived from Tyrosine-Containing Peptides and Simpler Hydroxyarenes Forming 4-(Diethoxyphosphinyl)phenylalanines and Diethyl Arylphosphonates," *J. Am. Chem. Soc. 109*:2831-2833, American Chemical Society (1987)	
	II	PITCHER, H.R., "Built-in Bypass," *Nature 429*:39, Nature Publishing Group (May 2004)	
	IJ	PREDVODITELEV, D.A., *et al.*, "Glycero-2-Hydroxymethylene Phosphates," *J. Org. Chem. USSR, A Translation of Zhur. Org. Khim. 13*:1489-1492, Plenum Publishing Corporation (1977)	
	IK	PREDVODITELEV, D.A., *et al.*, "Synthesis of Lipids and Their Models on the Basis of Glycerol Alkylene Phosphites. V. Cyclic Phosphatidylglycerol and Phosphatidylhydroxyhomocholine," *J. Org. Chem. USSR, A Translation of Zhur. Org. Khim. 17*:1156-1165, Plenum Publishing Corporation (1981)	
	IL	REDDY, K.R., *et al.*, "Stereoselective synthesis of nucleoside monophosphate HepDirect™ prodrugs," *Tetrahedron Lett. 46*:4321-4324, Elsevier Ltd. (2005)	
	IM	REDDY, M.R., *et al.*, "Development of a Quantum Mechanics-Based Free-Energy Perturbation Method: Use in the Calculation of Relative Solvation Free Energies," *J. Am. Chem. Soc. 126*:6224-6225, American Chemical Society (published online April 2004)	
	IN	REDMORE, D., "Phosphorus Derivatives of Nitrogen Heterocycles. 2. Pyridinephosphonic Acid Derivatives," *J. Org. Chem. 35*:4114-4117, American Chemical Society (1970)	
	IO	SARTILLO-PISCIL, F., *et al.*, "Fosfato-ésteres cíclicos diastereoisoméricos: 5-bromo-4-fenil-2-fenoxi-2-oxo-1,3,2-dioxafosforinanos, precursores de radicales libres alquilo β-fosfatoxi y generadores de radicales catiónicos en medio no oxidativo," *Revista de la Sociedad Química de México 46*:330-334, Sociedad Quimica de Mexico (2002)	

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	IP	SHAW, J.-P. and Cundy, K.C., "Biological Screens of PMEA Prodrugs," *Pharm. Res. 10*:S-294, Kluwer Academic Publishers B.V., Abstract No. PDD 7480 (1993)	
	IQ	SHIH, Y.-E., *et al.*, "Preparation and Structures of 2-Dimethylamino-4-phenyl-1,3,2-dioxaphosphorinane-2-oxides," *Bull. Inst. Chem., Academia Sinica 41*:9-16, Academia Sinica, Nankang, Taipei, Taiwan (1994)	
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	IS	THOMSON, W., *et al.*, "Synthesis, Bioactivation and Anti-HIV Activity of the Bis(4-acyloxybenzyl) and Mono(4-acyloxybenzyl) Esters of the 5'-monophosphate of AZT," *J. Chem. Soc. Perk. Trans. 1*,1239-1245, Royal Society of Chemistry (1993)	
	IT	VALENTINE, Jr., D., "Preparation of the Enantiomers of Compounds Containing Chiral Phosphorus Centers," *Asymmetric Synthesis* 4:263-312, Academic Press, Inc. (1984)	
	IU	VENOOK, A., "Treatment of Hepatocellular Carcinoma: Too Many Options?," *J. Clin. Oncol. 12*:1323-1334, American Society of Clinical Oncology (1994)	
	IV	VO-QUANG, Y., *et al.*, "(1-Amino-2-propenyl)phosphonic Acid, an Inhibitor of Alanine Racemase and D-Alanine:D-Alanine Ligase," *J. Med. Chem. 29*:579-581, American Chemical Society (1986)	
	IW	WAGNER, A., *et al.*, "Direct Conversion of Tetrahydropyranylated Alcohols to the Corresponding Bromides," *Tetrahedron Lett. 30*:557-558, Pergamon Press plc (1989)	
	IX	WALLACE, E.M., *et al.*, "Design and Synthesis of Potent, Selective Inhibitors of Endothelin-Converting Enzyme," *J. Med. Chem. 41*:1513-1523, American Chemical Society (1998)	
	IY	WALSH, E., *et al.*, "Phenoxymethylphosphonic Acids and Phosphonic Acid Ion-exchange Resins," *Phenoxymethylphosphonic Acid Ion-Exchange Resins 78*:4455-4458, American Chemical Society (1956)	

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	IZ	WATKINS, P., "Noninvasive tests of CYP3A enzymes," *Pharmacogenetics 4*:171-184, Lippincott Williams & Wilkins (1994)	
	JA	WEBER, G.F. and Waxman, D.J., "Activation of the Anti-cancer Drug Ifosphamide by Rat Liver Microsomal P450 Enzymes," *Biochem. Pharm. 45*:1685-1694, Pergamon Press Ltd. (1993)	
	JB	WEIBEL, M., *et al.*, "Potentiating Effect of {2-[2-[(2-Amino-1,6-Dihydro-6-Oxo-9H-Purin-9-yl)Methyl]-Phenyl] Ethenyl}-Phosphonic Acid (MDL 74,428), A Potent Inhibitor of Purine Nucleoside Phosphorylase, on the Antiretroviral Activities of 2',3'-Dideoxyinosine Combined to Ribavirin in Mice," *Biochem. Pharmacol. 48*:245-252, Elsevier Science Ltd. (1994)	
	JC	WILEMAN, T., *et al.*, "Receptor-mediated endocytosis," *Biochem. J. 232*:1-14, Portland Press (1985)	
	JD	YU, L. J., *et al.*, "In vivo Modulation of Alternative Pathways of P-450-Catalyzed Cyclophosphamide Metabolism: Impact on Pharmacokinetics and Antitumor Activity," *J. Pharmacol. Exp. Ther. 288*:928-937, The American Society for Pharmacology and Experimental Therapeutics (1999)	
	JE	ZON, G., "Cyclophosphamide Analogues" in *Progress in Medicinal Chemistry*, Ellis, G.P., *et al.*, eds., Elsevier Biomedical Press, Chapter 4, pp. 205-246 (1982)	
	JF	ZON, G., *et al.*, "NMR Spectroscopic Studies of Intermediary Metabolites of Cyclophosphamide. A Comprehensive Kinetic Analysis of the Interconversion of *cis*-and *trans*-4-Hydroxycyclophosphamide with Aldophosphamide and the Concomitant Partitioning of Aldophosphamide between Irreversible Fragmentation and Reversible Conjugation Pathways," *J. Med. Chem. 27*:466-485, American Chemical Society (1984)	
	JG	International Search Report for related International Application No. PCT/US03/34690, European Patent Office, Netherlands, mailed April 26, 2004	
	JH	Copy of Office Action for co-pending United States Application No. 10/698,924, Reddy, K.R., *et al.*, filed October 31, 2003, mailed June 22, 2005	

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